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<u>L3</u>	L2 same human	65	<u>L3</u>
<u>L2</u>	guanylyl cyclase	339	<u>L2</u>
<u>L1</u>	human guanylyl cyclase	0	<u>L1</u>

END OF SEARCH HISTORY

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Search Results - Record(s) 1 through 12 of 12 returned.

☐ 1. Document ID: US 20020155119 A1

L4: Entry 1 of 12

File: PGPB

Oct 24, 2002

PGPUB-DOCUMENT-NUMBER: 20020155119

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20020155119 A1

TITLE: Isolation and use of fetal urogenital sinus expressed sequences

PUBLICATION-DATE: October 24, 2002

INVENTOR-INFORMATION:

STATE COUNTRY RULE-47 NAME CITY Sikes, Robert A. US Gordonsville VA Lovingston VA US Chung, Leland W.K. Santa Monica CA US Kim, Jin Hee Fasciana, Claudia Rotterdam NLTrapman, Jan Mijnsheerenland NL

 $\text{US-CL-CURRENT: } \underline{424/185.1}; \ \underline{435/320.1}, \ \underline{435/325}, \ \underline{435/6}, \ \underline{435/69.1}, \ \underline{530/350}, \ \underline{536/23.5}$

Full Title Citation Front Review Classification Date Reference Sequences Attachments Claims KMIC Draw Desc Image

2. Document ID: US 6518013 B1

L4: Entry 2 of 12

File: USPT

Feb 11, 2003

US-PAT-NO: 6518013

DOCUMENT-IDENTIFIER: US 6518013 B1

TITLE: Methods for the inhibition of epstein-barr virus transmission employing

anti-viral peptides capable of abrogating viral fusion and transmission

Full Title Citation Front Review Classification Date Reference Sequences Attachments Claims KMC Draw Desc Image

3. Document ID: US 6500938 B1

L4: Entry 3 of 12

File: USPT

Dec 31, 2002

US-PAT-NO: 6500938

DOCUMENT-IDENTIFIER: US 6500938 B1

TITLE: Composition for the detection of signaling pathway gene expression

Full Title Citation Front Review Classification Date Reference Sequences Attachments Claims KMC Draw Desc Image

TITLE: Methods for inhibition of membrane fusion-associated events, including respiratory syncytial virus transmission Tol Title Cause front Review Customers Schemes Schemes Albertonis Cause Note Cause Cause 5. Document ID: US 6335170 B1 L4: Entry 5 of 12 File: USPT Jan 1, 2002 US-PAT-NO: 6335170 DOCUMENT-IDENTIFIER: US 6335170 B1 TITLE: Gene expression in bladder tumors 16. Document ID: US 6228983 B1 L4: Entry 6 of 12 File: USPT May 8, 2001 US-PAT-NO: 6228983 DOCUMENT-IDENTIFIER: US 6228983 B1 L4: Entry 6 of 12 File: USPT May 8, 2001 US-PAT-NO: 6228983 DOCUMENT-IDENTIFIER: US 6228983 B1 TITLE: Human respiratory syncytial virus peptides with antifusogenic and antiviral activities 7. Document ID: US 6093794 A L4: Entry 7 of 12 File: USPT Jul 25, 2000 US-PAT-NO: 6093794 DOCUMENT-IDENTIFIER: US 6093794 A L4: Entry 7 of 12 File: USPT Jul 25, 2000 US-PAT-NO: 6093794 DOCUMENT-IDENTIFIER: US 6093794 A TITLE: Isolated peptides derived from the Epstein-Barr virus containing fusion inhibitory domains	4. Document ID: US 64/	9055 BI	
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J. Document ID: US 6335170 B1 L4: Entry 5 of 12 File: USPT Jan 1, 2002 US-PAT-NO: 6335170 DOCUMENT-IDENTIFIER: US 6335170 B1 TITLE: Gene expression in bladder tumors Total Title Gaston Front Review Contribution Cate References Seasonces Albertients Staine Folice Danie Dess Incops G. Document ID: US 6228983 B1 L4: Entry 6 of 12 File: USPT May 8, 2001 US-PAT-NO: 6228983 DOCUMENT-IDENTIFIER: US 6228983 B1 ** See image for Certificate of Correction ** TITLE: Human respiratory syncytial virus peptides with antifusogenic and antiviral activities Total Folice Certificate Course Cate Reference Seasonces Albertier Links Date Date Dess Integral T. Document ID: US 6093794 A L4: Entry 7 of 12 File: USPT Jul 25, 2000 US-PAT-NO: 6093794 DOCUMENT-IDENTIFIER: US 6093794 A TITLE: Isolated peptides derived from the Epstein-Barr virus containing fusion inhibitory domains Total Title Certificate Front Review Constitution Date Reference Seasonces Albertieres Color Course Integral 8. Document ID: US 6068973 A			d events, including
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US-PAT-NO: 6068973

DOCUMENT-IDENTIFIER: US 6068973 A

TITLE: Methods for inhibition of membrane fusion-associated events, including influenza virus

Full Title Citation Front Review Classification Date Reference Sequences Attachments 9. Document ID: US 6060065 A May 9, 2000 L4: Entry 9 of 12 File: USPT US-PAT-NO: 6060065 DOCUMENT-IDENTIFIER: US 6060065 A TITLE: Compositions for inhibition of membrane fusion-associated events, including influenza virus transmission Full Title Citation Front Review Classification Date Reference Sequences Attachments KMC Draw Desc Image ☐ 10. Document ID: US 6054265 A File: USPT Apr 25, 2000 L4: Entry 10 of 12 US-PAT-NO: 6054265 DOCUMENT-IDENTIFIER: US 6054265 A TITLE: Screening assays for compounds that inhibit membrane fusion-associated events Full Title Citation Front Review Classification Date Reference Sequences Attachments KWIC Draw Desc Image ☐ 11. Document ID: US 6017536 A Jan 25, 2000 File: USPT L4: Entry 11 of 12 US-PAT-NO: 6017536 DOCUMENT-IDENTIFIER: US 6017536 A TITLE: Simian immunodeficiency virus peptides with antifusogenic and antiviral activities Full Title Citation Front Review Classification Date Reference Sequences Attachments KMC Draw Desc Image

☐ 12. Document ID: US 6013263 A

File: USPT L4: Entry 12 of 12

Jan 11, 2000

US-PAT-NO: 6013263

DOCUMENT-IDENTIFIER: US 6013263 A

TITLE: Measles virus peptides with antifusogenic and antiviral activities

Full Title Citation Front Review Classification Date Reference Sequences Attachments KMC Draw Desc Image Record List Display

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Previous Page Next Page







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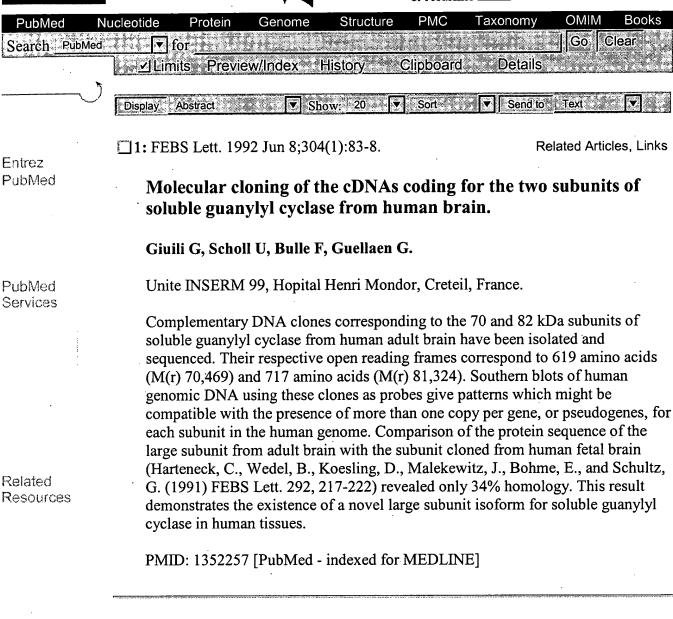
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INDEX 'ADISCTI, ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, AQUASCI, BIOBUSINESS, BIOCOMMERCE, BIOSIS, BIOTECHABS, BIOTECHDS, BIOTECHNO, CABA, CANCERLIT, CAPLUS, CEABA-VTB, CEN, CIN, CONFSCI, CROPB, CROPU, DDFB, DDFU, DGENE, DRUGB, DRUGLAUNCH, DRUGMONOG2, ...' ENTERED AT 11:50:47 ON 26 AUG 2003

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  34
       FILE AGRICOLA
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       FILE BIOTECHABS
 13
       FILE BIOTECHDS
 13
       FILE BIOTECHNO
 666
 83
       FILE CABA
163
       FILE CANCERLIT
2059
       FILE CAPLUS
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  28
       FILE CONFSCI
       FILE DDFB
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 183
       FILE DDFU
 130
       FILE DGENE
  3
       FILE DRUGB
   2
       FILE DRUGNL
 241
       FILE DRUGU
       FILE DRUGUPDATES
  2
       FILE EMBAL
 33
1772
       FILE EMBASE
1380
       FILE ESBIOBASE
 66
       FILE FEDRIP
  3
       FILE FROSTI
       FILE FSTA
  2
 508
       FILE GENBANK
       FILE IFIPAT
 34
       FILE JICST-EPLUS
 75
 498
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1858
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 11
       FILE PASCAL
 781
       FILE PHAR
 102
       FILE PHARMAML
  1
       FILE PHIN
  5
       FILE PROMT
 19
       FILE SCISEARCH
2613
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       FILE TOXCENTER
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       FILE USPAT2
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       FILE WPIDS
 30
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QUE GUANYLYL CYCLASE

FILE 'SCISEARCH, BIOSIS, CAPLUS, MEDLINE, EMBASE, ESBIOBASE, TOXCENTER, PASCAL, BIOTECHNO, LIFESCI, USPATFULL' ENTERED AT 11:52:11 ON 26 AUG 2003 3610 S L1 AND HUMAN 1370 S L2 AND (ISOLAT? OR CHARACTERI? OR PURIF?)

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L2

L3

ANSWER 61 OF 71 EMBASE COPYRIGHT 2003 ELSEVIER SCI. B.V. on STN

1999331767 EMBASE ACCESSION NUMBER:

Treatment of perioperative hypertension. TITLE:

Levy J.H. AUTHOR:

CORPORATE SOURCE: Dr. J.H. Levy, Department of Anesthesiology, Emory

University Hospital, 1364 Clifton Road, NE, Atlanta, GA

30322, United States

Anesthesiology Clinics of North America, (1999) 17/3 SOURCE:

> (567-579). Refs: 122

ISSN: 0889-8537 CODEN: ACNAEH

COUNTRY:

United States

Journal; General Review DOCUMENT TYPE: Internal Medicine FILE SEGMENT: 006

024 Anesthesiology

Drug Literature Index 037 Adverse Reactions Titles 038

LANGUAGE:

English

SUMMARY LANGUAGE: English Perioperative hypertension during cardiac or noncardiac surgery is a

unique clinical problem characterized by systemic vasoconstriction often with intravascular hypovolemia that usually requires acute short-term intravenous therapy. beta.-Adrenergic blockers are important first-line drugs for the patient with hypertension and tachycardia, although .beta.-blockers can have potential adverse side effects. The short-acting .beta.-blocker esmolol because of its titratability is a firstline .beta.-blocker for perioperative use. The CCBs represent important drugs with arterial vasodilating actions, and the new intravenous dihydropyridine compounds are especially promising because they have no negative inotropic effects or effects on atrioventricular node conduction. Nicardipine is the first intravenous dihydropyridine CCB currently available for perioperative hypertension in the United States, and clevidipine is currently under investigation. The following list summarizes therapeutic approaches to perioperative systemic hypertension:

.alpha.1- Adrenergic receptor blockade (phentolamine); ACE inhibition (enalaprilat); .beta.-Adrenergic blockade (esmolol, propranolol, metoprolol, atenolol); Calcium- channel blockade (nicardipine, isradipine, clevidipine); Dopamine-1 receptor stimulation (fenoldopam); Vascular guanylyl cyclase stimulation

(nitrovasodilators: nitroprusside, nitroglycerin); Vascular adenylyl cyclase stimulation (pulmonary hypertension: prostacyclin, prostaglandin E1).

L5 ANSWER 62 OF 71 Elsevier BIOBASE COPYRIGHT 2003 Elsevier Science B.V. on STN

ACCESSION NUMBER:

ESBIOBASE 1999063470

TITLE:

Muscarinic and .beta.-adrenergic regulation of heart rate, force of contraction and calcium current is preserved in mice lacking endothelial nitric oxide synthase

AUTHOR:

Vandecasteele G.; Eschenhagen T.; Scholz H.; Stein B.;

Verde I.; Fischmeister R.

CORPORATE SOURCE:

R. Fischmeister, Lab. Cardiol., Cell. et Moleculaire, INSERM U-446, Universite de Paris-Sud, F-92296

Chatenay-Malabry, France.

E-mail: Fisch@vjf.inserm.fr

Nature Medicine, (1999), 5/3 (331-334), 20 SOURCE:

reference(s)

CODEN: NAMEFI ISSN: 1078-8956

DOCUMENT TYPE:

Journal; Article

COUNTRY:

United States

LANGUAGE: SUMMARY LANGUAGE: English English

Nitric oxide (NO) is an ubiquitous signaling molecule produced from L-AB arginine by NO synthase (NOS). In the vasculature, NO mediates parasympathetic endothelium-dependent vasodilation. NO may also mediate the parasympathetic control of myocardial function. This is supported by the observations that NOS3, the endothelial constitutive NOS, is expressed in normal cardiac myocytes from rodents and human, and NOS and/or guanylyl cyclase inhibitors antagonize the effect of muscarinic agonists on heart rate, atrio-ventricular conduction, contractility and L-type calcium current. Here we examine the autonomic regulation of the heart in genetically engineered mice deficient in NOS3 (NOS3-KO) (ref. 8). We show that the chronotropic and inotropic responses to both .beta.-adrenergic and muscarinic agonists were unaltered in isolated cardiac tissue preparations from NOS3-KO mice, although these mice have a defective parasympathetic regulation of vascular tone. Similarly, .beta.-adrenergic stimulation and muscarinic inhibition of the calcium current did not differ in cardiac myocytes from NOS3-KO mice and those from wild-type mice. RT-PCR did not demonstrate upregulation of other NOS isoforms. Similarly, G(i)/G(o) proteins and muscarinic receptor density were unaltered. These data refute the idea that NOS3 is obligatory for the normal autonomic control of cardiac muscle function.

ANSWER 63 OF 71 SCISEARCH COPYRIGHT 2003 THOMSON ISI on STN DUPLICATE 6

1998:102849 SCISEARCH ACCESSION NUMBER:

THE GENUINE ARTICLE: YT669

Characterization of NS 2028 as a specific TITLE:

inhibitor of soluble guanylyl cyclase

Olesen S P (Reprint); Drejer J; Axelsson O; Moldt P; Bang AUTHOR:

L; NielsenKudsk J E; Busse R; Mulsch A

NEUROSEARCH, 26B SMEDLAND, DK-2600 GLOSTRUP, DENMARK CORPORATE SOURCE:

(Reprint); RIGSHOSP, DEPT MED B, DK-2100 COPENHAGEN O, DENMARK; UNIV FRANKFURT KLINIKUM, ZENTRUM PHYSIOL, D-60590

FRANKFURT, GERMANY

COUNTRY OF AUTHOR:

DENMARK; GERMANY

SOURCE:

BRITISH JOURNAL OF PHARMACOLOGY, (JAN 1998) Vol. 123, No.

2, pp. 299-309.

Publisher: STOCKTON PRESS, HOUNDMILLS, BASINGSTOKE,

HAMPSHIRE, ENGLAND RG21 6XS.

ISSN: 0007-1188.

DOCUMENT TYPE: FILE SEGMENT:

Article; Journal

LANGUAGE:

LIFE English

REFERENCE COUNT:

ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS AB

1 The haeme-containing soluble guanylyl cyclase (alpha(1) beta(1)-heterodimer) is a major intracellular receptor and effector for nitric oxide (NO) and carbon monoxide (GO) and mediates many of their biological actions by increasing cyclic GMP. We have synthesized new oxadiazolo-benz-oxazins and have assessed their inhibitory actions on guanylyl cyclase activity in vitro, on the formation of cyclic GMP in cultured cells and on the NO-dependent relaxation of vascular and non-vascular smooth muscle.

2 Soluble guanylyl cyclase, purified to homogeneity from bovine lung, was inhibited by 4H-8-bromo-1,2,4oxadiazolo(3,4-d)benz(b)(1,4)oxazin-1-one (NS 2028) in a concentration-dependent and irreversible manner (IC50 30 nM for basal and 200 nM for NO-stimulated enzyme activity). Evaluation of the inhibition kinetics according to Kitz & Wilson yielded a value of 8 nM for K-i, the equilibrium constant describing the initial reversible reaction between inhibitor and enzyme, and 0.2 min(-1) for the rate constant k3 of the subsequent irreversible inhibition. Inhibition was accompanied by a shift in the soret absorption maximum of the enzyme's haem cofactor from 430 to 390 nm.

3 S-nitroso-glutathione-enhanced soluble guanylyl

cyclase activity in homogenates of mouse cerebellum was inhibited by NS 2028 (IC50 17 nM) and by 17 structural analogues in a similar manner, albeit with different potency, depending on the type of substitution at positions 1, 7 and 8 of the benzoxazin structure. Small electronegative ligands such as Br and Cl at position 7 or 8 increased and substitution of the oxygen at position 1 by -S-,- NH- or -CH2- decreased the inhibition.

4 In tissue slices prepared from mouse cerebellum, neuronal NO synthase-dependent activation of soluble guanylyl cyclase by the glutamate receptor agonist N-methyl-D-aspartate was inhibited by NS 2028 (IC50 20 nM) and by two of its analogues. Similarly, 3-morpholino-sydnonimine (SIN-1)-elicited formation of cyclic GMP in human cultured umbilical vein endothelial cells was inhibited by NS 2028 (IC50 30 nM).

5 In prostaglandin F-2 alpha-constricted, endothelium-intact porcine coronary arteries NS 2028 elicited a concentration-dependent increase (65%) in contractile tone (EC50 170 nM), which was abolished by removal of the endothelium. NS 2028 (1 mu M) suppressed the relaxant response to nitroglycerin from 88.3+/-2.1 to 26.8+/-6.4% and induced a 9 fold rightward shift (EC50 15 mu M) of the concentration-relaxation response curve to nitroglycerin. It abolished the relaxation to sodium nitroprusside (1 mu M), bur did not affect the vasorelaxation to the K-ATPchannel opener cromakalim. Approximately 50% of the relaxant response to sodium nitroprusside was recovered after 2 h washout of NS 2028.

6 In phenylephrine-preconstricted, endothelium-denuded aorta of the rabbit NS 2028 (1 mu M) did not affect relaxant responses to atrial natriuretic factor, an activator of particulate guanylyl cyclase, or forskolin, an activator of adenylyl cyclase.

7 NO-dependent relaxant responses in non-vascular smooth muscle were also inhibited by NS 2028. The nitroglycerin-induced relaxation of guinea-pig trachea preconstricted by histamine was fully inhibited by NS 2028 (1 mu M), whereas the relaxations to terbutaline, theophylline and vasoactive intestinal polypeptide (VIP) were not affected. The relaxant responses to electrical field stimulation of non-adrenergic, non-cholinergic nerves in the same tissue were attenuated by 50% in the presence of NS 2028 (1 mu M)

8 NS 2028 and its analogues, one of which is the previously characterized 1H-[1,2,4]oxadiazolo[4,3, a]quinoxalin-1-one (ODQ), appear to be potent and specific inhibitors of soluble guanylyl cyclase present in various cell types. Oxidation and/or a change in the coordination of the haeme-iron of guanylyl cyclase is a likely inhibitory mechanism.

ANSWER 64 OF 71 SCISEARCH COPYRIGHT 2003 THOMSON ISI on STN DUPLICATE 7 L_5 ACCESSION NUMBER: 1998:818230 SCISEARCH

THE GENUINE ARTICLE: 130KM

Functional properties of a naturally occurring isoform of TITLE:

soluble guanylyl cyclase

Russwurm M; Behrends S; Harteneck C; Koesling D (Reprint) AUTHOR:

FREE UNIV BERLIN, INST PHARMAKOL, THIELALLEE 69-73,

D-14195 BERLIN, GERMANY (Reprint); FREE UNIV BERLIN, INST

PHARMAKOL, D-14195 BERLIN, GERMANY

COUNTRY OF AUTHOR:

CORPORATE SOURCE:

GERMANY

SOURCE:

BIOCHEMICAL JOURNAL, (1 OCT 1998) Vol. 335, Part 1, pp.

125-130.

Publisher: PORTLAND PRESS, 59 PORTLAND PLACE, LONDON W1N

3AJ, ENGLAND. ISSN: 0264-6021.

DOCUMENT TYPE:

Article; Journal LIFE

FILE SEGMENT: LANGUAGE:

AB

English

REFERENCE COUNT:

ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS

Soluble guanylyl cyclase (sGC), the target enzyme

of the signalling molecule NO, contains one prosthetic haem group and consists of an alpha and a beta subunit. So far, only the alpha (1) beta(1) heterodimer has been shown to exist in different cells and tissues, and most biochemical studies of sGC have been performed with the alpha(1)beta(1) heterodimer. Here we demonstrate for the first time the natural occurrence of the a, subunit on the protein lever. The alpha(2) subunit co-precipitated with the beta(1) subunit from human placenta, showing the existence of the alpha(2)beta (1) isoform in vivo. The new enzyme was expressed in and purified from cells from the Spodoptera frugiperda ovary cell line Sf 9. Spectral analysis showed that the alpha(2)beta(1) heterodimer contains a prosthetic haem group revealing the same characteristics as the haem in the alpha(1) beta(1) form. The kinetic properties of both isoforms and sensitivity towards NO were indistinguishable. H-1-[1,2,4]oxadiazolo[4,3-a]quinoxalin-1-one (ODQ), a selective inhibitor of sGC, abolished NO-stimulated activity of both heterodimers. The new NO-independent activator, 3-(5'-hydroxymethyl-2'-furyl)-1-benzyl indazole (YC-1), increased the maximal NO-stimulated activity of the new isoform, caused a leftward-shift in the NO concentration-response curve and turned CO into an effective activator, as it did for the alpha(1) beta(1) heterodimer (200-fold activation). In summary, the differences in primary structure of both a subunits are contrasted by their functional similarity. Further studies will be needed to elucidate the physiological purpose of the new isoform.

ANSWER 65 OF 71 USPATFULL on STN

ACCESSION NUMBER:

97:114932 USPATFULL

TITLE:

INVENTOR(S):

Suppression of nitric oxide production by osteopontin Denhardt, David T., Bridgewater, NJ, United States Hwang, Shiaw-Min, Piscataway, NJ, United States Heck, Diane Elaine, Rumson, NJ, United States

Lopez, Cecilia Ang, North Brunswick, NJ, United States Laskin, Debra L., Basking Ridge, NJ, United States Laskin, Jeffrey D., Piscataway, NJ, United States Rutgers University, Piscataway, NJ, United States (U.S.

PATENT ASSIGNEE(S):

corporation)

University of Medicine & Dentistry of NJ, Newark, NJ,

United States (U.S. corporation)

	NUMBER	KIND	DATE	
			10071000	
PATENT INFORMATION:	US 5695761 ·		19971209	
APPLICATION INFO.:	US 1993-173116		19931223	(8)
DOCUMENT TYPE:	Utility			
FILE SEGMENT:	Granted			
PRIMARY EXAMINER:	Hutzell, Paula K.			
ASSISTANT EXAMINER:	Minnifield, N. M.			
LEGAL REPRESENTATIVE:	Klauber & Jackson			
NUMBER OF CLAIMS:	18			
EXEMPLARY CLAIM:	1			

19 Drawing Figure(s); 7 Drawing Page(s) NUMBER OF DRAWINGS:

LINE COUNT: 1552

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention relates to compositions and methods for inhibition of the vasoactive and signal transduction agent nitric oxide (NO), and to therapeutic treatment of diseases or disorders that involve inappropriate or detrimental NO activity. The invention particularly relates to modulation of kidney function. In specific embodiments, osteopontin and a 20-amino acid fragment of osteopontin that contains an Arg-Gly-Asp sequence suppress expression of inducible NO synthase mRNA, and osteopontin suppresses the activity of constitutive NO synthase.

ANSWER 66 OF 71 USPATFULL on STN

97:78170 USPATFULL ACCESSION NUMBER:

Compositions and methods for cancer immunotherapy TITLE: Barber, Jack R., San Diego, CA, United States INVENTOR(S): Jolly, Douglas J., Leucadia, CA, United States Respess, James G., San Diego, CA, United States

Chiron Viagene, Inc., San Diego, CA, United States PATENT ASSIGNEE(S):

(U.S. corporation)

KIND NUMBER DATE ______

PATENT INFORMATION:

19970902 US 5662896 US 1993-32846 19930317 (8)

APPLICATION INFO .: RELATED APPLN. INFO.:

Continuation-in-part of Ser. No. US 1992-965084, filed on 22 Oct 1992, now abandoned which is a continuation of Ser. No. US 1990-586603, filed on 21 Sep 1990, now abandoned which is a continuation-in-part of Ser. No.

US 1990-565606, filed on 10 Aug 1990, now abandoned which is a continuation-in-part of Ser. No. US

1989-395932, filed on 18 Aug 1989, now abandoned which is a continuation-in-part of Ser. No. US 1988-170515,

filed on 21 Mar 1988, now abandoned

DOCUMENT TYPE: FILE SEGMENT:

Utility Granted

PRIMARY EXAMINER: ASSISTANT EXAMINER:

Fleisher, Mindy Railey, II, Johnny F.

LEGAL REPRESENTATIVE:

Seed & Berry, Kruse, Norman J., Blackburn, Robert P.

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS:

34 Drawing Figure(s); 22 Drawing Page(s)

LINE COUNT:

2662

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention provides methods for inhibiting the growth of selected tumors utilizing recombinant viral vectors. Briefly, within one aspect of the present invention, a method for inhibiting the growth of a selected tumor is provided comprising the step of directly administering to a warm-blooded animal a vector construct which directs the expression of at least one anti-tumor agent, such that the growth of said tumor is inhibited. Representative examples of anti-tumor agents include immune activators and tumor proliferation inhibitors.

ANSWER 67 OF 71 USPATFULL on STN

ACCESSION NUMBER:

97:20543 USPATFULL

TITLE:

Use of .alpha..sub.1A -selective adrenoceptor agonists

for the treatment of urinary incontinence

INVENTOR(S):

Craig, Douglas A., Fair Lawn, NJ, United States Forray, Carlos C., Paramus, NJ, United States Gluchowski, Charles, Wayne, NJ, United States Branchek, Theresa A., Teaneck, NJ, United States Synaptic Pharmaceutical Corporation, Paramus, NJ,

PATENT ASSIGNEE(S):

United States (U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: . APPLICATION INFO.:

US 5610174 19970311 US 1995-459410 19950602 (8)

DOCUMENT TYPE: FILE SEGMENT:

Utility Granted

PRIMARY EXAMINER: ASSISTANT EXAMINER:

Jordan, Kimberly Jarvis, William R. A.

LEGAL REPRESENTATIVE:

White, John P.

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

3 1

NUMBER OF DRAWINGS:

16 Drawing Figure(s); 6 Drawing Page(s)

LINE COUNT: 1626

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention provides a method of treating urinary incontinence in a subject which comprises administering to the subject a therapeutically effective amount of a compound having the following structure: ##STR1## wherein each of the substituents for the compound is as defined in the specification.

L5 ANSWER 68 OF 71 LIFESCI COPYRIGHT 2003 CSA on STN

ACCESSION NUMBER:

CORPORATE SOURCE:

96:100765 LIFESCI

TITLE:

Assignment of GUCIA2, the gene coding for the alpha 2

subunit of soluble guanylyl cyclase, to position 11q21-q22 on human chromosome 11

AUTHOR:

Yu, F.; Warburton, D.; Wellington, S.; Danziger, R.S.* Div. Cardiology, Coll. Physicians and Surgeons Columbia

Univ., 630 W. 168th St., New York, NY 10032, USA

SOURCE:

GENOMICS, (1996) vol. 33, no. 2, pp. 334-336.

ISSN: 0888-7543.

DOCUMENT TYPE:

Journal

FILE SEGMENT:

English

LANGUAGE:

Soluble guanylyl cyclases, which are activated by

nitric oxide (NO), are obligate heterodimers (alpha / beta) with an associated heme group for binding NO. Two isoforms of each subunit, i.e.,

alpha 1, alpha 2, beta 1, and

beta 2, have been characterized. The alpha 1 (82 kDa) and beta 1 (73 kDa) subunits were first

purified as a heterodimer from bovine lung and subsequently cloned from rat, bovine, and human lungs. A second isoform of the beta

subunit (beta 2) was cloned from the rat kidney. More recently, a second

isoform of the alpha subunit (alpha 2) was cloned from human

brain. The alpha subunits have been shown to be interchangeable such

that heterodimers consisting of alpha 1/ beta

1 and alpha 2/ beta 1 subunits are active,

but not those of alpha 1/ alpha 2. Furthermore, the

available data suggest that there is tissue-specific expression of the subunit isoforms, i.e., lung tissue contains alpha 1

and beta 1, the cortical collecting duct of the

kidney contains alpha 1 and beta 2, and the

alpha 1, beta 1, and beta 2

subunit isoforms predominate in the renal vasculature. The genes coding

for the alpha 1 and beta 1

subunits of soluble guanylyl cyclase (GUCIA3, GUCIB3)

have been colocalized to the same human chromosome region, i.e.,

chromosome 4 at q31.3-q33. In the current study, we have shown that the gene for the alpha 2 subunit, GUCIA2, is located at 11q21-q22 on

human chromosome 11.

L5 ANSWER 69 OF 71 SCISEARCH COPYRIGHT 2003 THOMSON ISI ON STN DUPLICATE 8 ACCESSION NUMBER: 95:430966 SCISEARCH

THE GENUINE ARTICLE: RE666

TITLE: 2

2 DROSOPHILA GENES THAT ENCODE THE ALPHA-SUBUNIT AND

BETA-SUBUNIT OF THE BRAIN SOLUBLE GUANYLYL

CYCLASE

AUTHOR:

SHAH S; HYDE D R (Reprint)

CORPORATE SOURCE: UNIV NOTRE DAME, DEPT BIOL SCI, NOTRE DAME, IN, 46556

(Reprint); UNIV NOTRE DAME, DEPT BIOL SCI, NOTRE DAME, IN,

46556

COUNTRY OF AUTHOR:

USA

SOURCE:

JOURNAL OF BIOLOGICAL CHEMISTRY, (23 JUN 1995) Vol. 270,

No. 25, pp. 15368-15376.

ISSN: 0021-9258.

DOCUMENT TYPE:

Article; Journal

FILE SEGMENT:

LIFE

ENGLISH LANGUAGE:

REFERENCE COUNT: 38

ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS

We identified two Drosophila genes (dgc alpha 1 and AB

dgc beta 1) that encode the soluble guanylyl

cyclase alpha and beta subunits, respectively. The putative Dgc

alpha 1 protein is 76 kDa, has 35% amino acid identity

with previously isolated alpha subunits, and was immunolocalized to the adult retina, to the optic lobes, and throughout the brain

neuropil. The Dgc beta 1 protein is 86 kDa and

exhibits 59% amino acid identity with the rat beta 1

protein. However, the Dgc beta 1 protein has an

additional 118 amino acids inserted near the amino terminus, which makes

it significantly larger than the rat beta 1. The Dgc beta 1 protein was immunolocalized to the optic lobes

and throughout the brain neuropil, with no detectable expression in the

retina. The Dgc alpha 1 and Dgc beta

1 cDNAs were stably transfected into human kidney 293

cells. Expression of the individual subunits and mixing of the individually expressed subunits failed to generate significant

guanylyl cyclase activity. Only coexpression of the subunits resulted in significant guanylyl cyclase

activity. Our results indicate that Dgc alpha 1 and Dgc beta 1 are soluble guanylyl

cyclase alpha and beta subunits that are capable of forming a

functional guanylyl cyclase heterodimer.

ANSWER 70 OF 71 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC. on STN

DUPLICATE 9

ACCESSION NUMBER: 1993:504357 BIOSIS

DOCUMENT NUMBER: PREV199396128364

Characterization of soluble guanylyl TITLE:

cyclase in transformed human non-pigmented epithelial cells.

Danziger, Robert S. (1); Star, Robert A. (1); Matsumoto, AUTHOR (S):

Shun; Coca-Prados, Miguel; Desantis, Louis; Pang, Iok-Hou

CORPORATE SOURCE: (1) Dep. Internal Med., Univ. Texas Southeastern Med.

Cent., Dallas, TX USA

Biochemical and Biophysical Research Communications, (1993) SOURCE:

Vol. 195, No. 2, pp. 958-962.

ISSN: 0006-291X.

DOCUMENT TYPE:

LANGUAGE:

Article English

Topical application of nitro vasodilators, such as sodium nitroprusside,

reduces intraocular pressure. In brain and blood vessels, nitro

vasodilators activate soluble guanylyl cyclases,

producing cGMP. The location and molecular identity of ocular

guanylyl cyclases are unknown: We studied transformed

human non-pigmented ciliary epithelial cells, whose parental cells are responsible for the production of aqueous humor. Sodium nitroprusside increased cGMP 40 to 60 fold in a time and concentration dependent manner (EC-50 40 to 100 mu-M). Methylene blue inhibited this effect (IC-50 0.6 mu-M in the presence of 100 mu-M sodium nitroprusside). We also detected

mRNA for the alpha-1 and beta-1,

but not the beta-2, subunit isoforms of soluble guanylyl cyclase in these cells by Northern blotting. We conclude that

transformed non-pigmented epithelial cells contain an active heterodimeric soluble guanylyl cyclase composed of at least

alpha-1 and beta-1 subunits.

L5 ANSWER 71 OF 71 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC. on STN

DUPLICATE 10

ACCESSION NUMBER: 1993:70686 BIOSIS DOCUMENT NUMBER: PREV199395035186

TITLE:

Characterization of soluble platelet

guanylyl cyclase with peptide antibodies.

AUTHOR(S):

Guthmann, Florian; Mayer, Bernd; Koesling, Doris; Kukovetz,

Walter R.; Boehme, Eycke (1)

CORPORATE SOURCE:

(1) Institut fuer Pharmakologie, Freie Universitaet Berlin,

Thielalle 67-73, W-1000 Berlin 33 Germany

SOURCE:

Naunyn-Schmiedeberg's Archives of Pharmacology, (1992) Vol.

346, No. 5, pp. 537-541.

ISSN: 0028-1298.

DOCUMENT TYPE:

Article English

LANGUAGE:

AB

Soluble guanylyl cyclase partially purified

from bovine and human platelets was characterized with

antibodies raised against synthetic peptides corresponding to different

sequences of the alpha-1- and beta-1

-subunits of the bovine lung enzyme. On immunoblots, the platelet

guanylyl cyclase was recognized by the four antisera

used, with the exception of an antiserum against the C-terminus of the

beta-1-subunit which did not react with the

human platelet but with the bovine platelet beta-

1-subunit. Furthermore the human platelet beta

-1-subunit exhibited a slightly lower molecular mass than the

bovine protein. The C-terminal antibodies precipitated native platelet and lung quanyly cyclase activity. In contrast an antibody against a peptide out of the putative catalytic domain, which is highly conserved between

all quanylyl cyclases sequenced so far, did not

precipitate native guanylyl cyclase, although it

recognized both subunits on immunoblots, suggesting that the respective

amino acid sequence is located in an inner site of the protein.